- 1. A non-naturally occurring compound comprising at least one methyl sulfide or methyl sulfoxide moiety, the compound being a substrate for at least one MsrA enzyme and at least one MsrB enzyme, or a pharmaceutically acceptable salt thereof.
- 2. The compound of claim 1, having formula 2, or a pharmaceutically acceptable salt thereof:

$$R_7$$
 $R_7$ 
 $R_4$ 
 $R_7$ 
 $R_4$ 
 $R_6$ 
 $R_7$ 
 $R_4$ 
 $R_7$ 
 $R_8$ 

wherein:

 $R_1$  is CH of either R or S configuration;  $R_2$  is a normal or branched alkyl or fluoroalkyl group having 1 to 6 carbons;  $R_3$  is methyl or ethyl or a fluorinated derivative thereof;  $R_4$  is a hydrogen or a normal or branched alkyl group having 1 to 6 carbons;  $R_5$  is a CH of either R or S configuration;  $R_6$  is a hydrogen or a normal or branched alkyl or fluoroalkyl group having 1 to 6 carbons;  $R_7$  is a nitrogen with substituent  $R_4$  as defined herein, a CH of either R or S configuration, or a normal or branched alkyl or fluoroalkyl group having 1 to 6 carbons; and S is either S or S in any oxidation state.

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3. The compound of claim 1, having formula 3, or a pharmaceutically acceptable salt thereof.:

wherein:

R<sub>1</sub> is CH of either R or S configuration; R<sub>2</sub> is a normal or branched alkyl or fluoroalkyl group having 1 to 6 carbons; R<sub>3</sub> is methyl or ethyl or a fluorinated derivative thereof; R<sub>4</sub> is a hydrogen or a normal or branched alkyl group having 1 to 6 carbons; R<sub>5</sub> is a CH of either R or S configuration; R<sub>6</sub> is a hydrogen or a normal or branched alkyl or fluoroalkyl group having 1 to 6 carbons; and X is either S or S in any oxidation state.

## 4.

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The compound of claim 1, having formula 4, or a pharmaceutically acceptable salt thereof:

$$R_{5}$$
  $R_{2}$   $R_{2}$   $R_{2}$   $R_{3}$   $R_{4}$   $R_{6}$   $R_{6}$   $R_{7}$   $R_{1}$   $R_{4}$   $R_{6}$   $R_{7}$   $R_{8}$ 

10 wherein:

> R<sub>1</sub> is CH of either R or S configuration; R<sub>2</sub> is a normal or branched alkyl or fluoroalkyl group having 1 to 6 carbons; R<sub>3</sub> is methyl or ethyl or a fluorinated derivative thereof; R<sub>4</sub> is a hydrogen or a normal or branched alkyl group having 1 to 6 carbons; R<sub>5</sub> is a CH of either R or S configuration; R<sub>6</sub> is a hydrogen or a normal or branched alkyl or fluoroalkyl group having 1 to 6 carbons; and X is either S or Se in any oxidation state.

> 5. The compound of claim 1, having formula 5, or a pharmaceutically acceptable salt thereof:

$$R_{6}$$
 $CH_{3}$ 
 $R_{7}$ 
 $R_{4}$ 
 $R_{4}$ 
 $R_{4}$ 
 $R_{4}$ 
 $R_{4}$ 
 $R_{5}$ 
 $R_{7}$ 
 $R_{1}$ 
 $R_{2}$ 
 $R_{3}$ 

20 wherein:

> R<sub>1</sub> is CH of either R or S configuration; R<sub>2</sub> is a normal or branched alkyl or fluoroalkyl group having 1 to 6 carbons; R<sub>3</sub> is methyl or ethyl or a fluorinated derivative thereof; R<sub>4</sub> is a hydrogen or a normal or branched alkyl group having 1 to 6 carbons; and X is either S or Se in any oxidation state.

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6. The compound of claim 1, having formula 2a, or a pharmaceutically acceptable salt thereof:

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7. The compound of claim 1, having formula 3a, or a pharmaceutically acceptable salt thereof.

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8. The compound of claim 1, having formula 4a, or a pharmaceutically acceptable salt thereof.

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9. The compound of claim 1, having formula 5a, or a pharmaceutically acceptable salt thereof.

10. A non-naturally occurring compound comprising at least one methyl sulfide or methyl sulfoxide moiety, the compound being a substrate for at least one Msr enzyme, said compound having a backbone not based on sulindac (1(Z)-5-fluoro-2-methyl-1[[4-(methylsulfinyl)phenyl)methylene]-1H-indenyl-3-acetic acid).

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11. The compound of claim 10 having formula 6, or a pharmaceutically acceptable salt thereof:

wherein:

15 the aromatic ring includes one or more nitrogen atoms; the aromatic carboxyl group is oriented ortho, meta, or para to the methionine-based moiety; R<sub>1</sub> is CH of either R or S configuration; R<sub>2</sub> is a normal or branched alkyl or fluoroalkyl group having 1 to 6 carbons; R<sub>3</sub> is methyl or ethyl or a fluorinated derivative thereof; R<sub>4</sub> is a hydrogen or a normal or branched alkyl group having 1 to 6 carbons; R<sub>5</sub> is a nitrogen with substituent R<sub>4</sub> as defined herein, an oxygen, or a sulfur; and X is S or Se in any oxidation state.

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12. The compound of claim 10 having formula 6a, or a pharmaceutically acceptable salt thereof:

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13. The compound of claim 10 having formula 7, or a pharmaceutically acceptable salt thereof:

OH 
$$R_2$$
  $R_2$   $R_3$   $R_4$   $R_4$   $R_5$   $R_4$   $R_6$   $R_7$   $R_8$ 

wherein:

both aromatic rings comprises one or more nitrogen atoms; the aromatic carboxyl group is oriented *ortho*, *meta*, or *para* to the aniline nitrogen;  $R_1$  is CH of either R or S configuration;  $R_2$  is normal or branched alkyl or fluoroalkyl group having 1 to 6 carbons;  $R_3$  is methyl or ethyl or a fluorinated derivative thereof;  $R_4$  is a hydrogen or a normal or branched alkyl group having 1 to 6 carbons; and X is S or S in any oxidation state.

14. The compound of claim 10 having formula 7a, or a pharmaceutically acceptable salt thereof:

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15. The compound of claim 10 having formula 8, or a pharmaceutically acceptable salt thereof:

$$\begin{array}{c|c}
 & O & R_2 \\
\hline
 & N - R_1 \\
 & O \\$$

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wherein:

the aromatic ring comprises one or more nitrogen atoms; the sec-butyl group is oriented ortho, meta, or para to the methionine-based moiety;  $R_1$  is CH of either R or S configuration;  $R_2$  is a normal or branched alkyl or fluoroalkyl group having 1 to 6 carbons;  $R_3$  is methyl or ethyl or a fluorinated derivative thereof;  $R_4$  is a hydrogen or a normal or branched alkyl group

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- having 1 to 6 carbons;  $R_5$  is a CH of either R or S configuration; X is either S or Se in any oxidation state.
  - 16. The compound of claim 10 having formula 8a, or a pharmaceutically acceptable salt thereof.

17. The compound of claim 10 having formula 9, or a pharmaceutically acceptable salt thereof:

$$R_{5}$$
 $R_{2}$ 
 $R_{2}$ 
 $R_{2}$ 
 $R_{3}$ 
 $R_{4}$ 
 $R_{4}$ 
 $R_{5}$ 
 $R_{7}$ 
 $R_{7}$ 

wherein:

Groups  $R_1$ ,  $R_2$ ,  $R_3$ ,  $R_4$ ,  $R_5$ ,  $R_6$ ,  $R_7$  and X in general structure 9 are defined as follows:  $R_1$  is CH of either R or S configuration;  $R_2$  is a normal or branched alkyl or fluoroalkyl group having 1 to 6 carbons;  $R_3$  is methyl or ethyl or a fluorinated derivative thereof;  $R_4$  is a hydrogen or a normal or branched alkyl group having 1 to 6 carbons;  $R_5$  is a CH of either R or S configuration;  $R_6$  is a hydrogen or a normal or branched alkyl or fluoroalkyl group having 1 to 6 carbons;  $R_7$  is any halogen oriented *ortho*, *meta*, or *para* to the carbonyl group, and X is S or Se in any oxidation state.

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18. The compound of claim 10 having formula 9a, or a pharmaceutically acceptable salt thereof:

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19. The compound of claim 10 having formula 10, or a pharmaceutically acceptable salt thereof:

wherein:

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the lactone ring is oriented *ortho*, *meta*, or *para* to the sulfonyl group;  $R_1$  is CH of either R or S configuration;  $R_2$  is a normal or branched alkyl or fluoroalkyl group having 1 to 6 carbons;  $R_3$  is methyl or ethyl or a fluorinated derivative thereof;  $R_4$  is a hydrogen or a normal or branched alkyl group having 1 to 6 carbons; X is S or Se in any oxidation state; S0 are phenyl, alkyl, halogen substituted phenyl, or heteroaromatic compound.

20. The compound of claim 10 having formula 10a, or a pharmaceutically acceptable salt thereof:

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21. A composition comprising the compound of claim 1 and a pharmaceutically acceptable carrier.

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22. A composition comprising the compound of claim 10 and a pharmaceutically acceptable carrier.

23. A method for reducing, preventing or reversing oxidative damage in a cell, the method comprising the steps of:

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- (a) providing a non-naturally occurring compound comprising at least one methyl sulfide or methyl sulfoxide moiety, the compound being a substrate for at least one Msr enzyme;
- (b) providing a cell expressing at least one Msr enzyme, said cell comprising or being exposed to reactive oxygen species; and

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- (c) contacting the cell with an amount of the compound sufficient to reduce, prevent, or reverse oxidative damage in the cell by said reactive oxygen species.
  - 24. The method of claim 23, wherein the cell is within an animal subject.

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25. The method of claim 23, wherein the animal subject has a condition or disorder associated with oxidative damage.

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27. The method of claim 23, wherein the condition is age-related.

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The method of claim 23, wherein the disorder involves degeneration of a

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28. A method for extending the lifespan of an animal comprising administering to the animal a therapeutically effective amount of a non-naturally occurring compound comprising at least one methyl sulfide or methyl sulfoxide moiety, the compound being a substrate for at least one Msr enzyme.

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